CT RLD INTELLECTUAL PROPERTY ORGANIZATIO.



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

			MINENII (FCI)
(51) International Patent Classification 7:		(11) International Publication Number:	WO 00/28081
C12Q 1/68	A3		
		(43) International Publication Date:	18 May 2000 (18.05.00)
(21) International Application Number: PCT/IB99/01958 (22) International Filing Date: 9 November 1999 (09.11.99)		(81) Designated States: AU, CA, JP, US, European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE).	
(71) Applicant (for all designated States except US): ME N.V. [BE/BE]; Onafhankelijkheidslaan 38, B-90 (BE). (72) Inventors; and (75) Inventors/Applicants (for US only): ZABEAU (BE/BE]; Onafhankelijkheidslaan 38, B-9000 Ge STANSSENS, Patrick [BE/BE]; Constant Permeke B-9830 St. Martens Laten (BE). (74) Agent: DE CLERCQ, Ann; Ann De Clercq & Co B Brandstraat 100, B-9830 Sint-Martens-Latem (BE)	THEXIOO Ge	(88) Date of publication of the internation of	the receipt of amendments.

(54) Title: RESTRICTED AMPLICON ANALYSIS

(57) Abstract

The present invention generally provides a method which facilitates the detection of polymorphisms (or mutations). The method is directed to the analysis of so-called endonuclease site polymorphisms (ESPs) that result in the gain or loss of a restriction endonuclease site. In essence, the ESP is probed with the restriction endonuclease reagent prior to amplification, whereby amplification is prevented and consequently no signal is observed when cleavage takes place. Unambiguous allele calling is performed by comparing the signals obtained with and without cleavage with the restriction endonuclease reagent. The method is particularly useful for multiplex genotyping, involving the parallel analysis of large numbers of single nucleotide polymorphisms. Preferred methods for detecting the amplicons involve hybridization to an arrayed or otherwise identifiable set of cognate probe fragments or oligonucleotides.